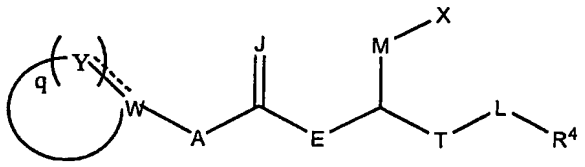
Claims

We claim:

1. A pharmaceutical composition comprising a compound of the structure (I)



(I)

wherein Y, at each occurrence, is independently selected from the group consisting of C(O), N, CR^1, $C(R^2)(R^3)$, NR^5, CH, O and S;

q is an integer of from 3 to 10;

A is selected from the group consisting of O, S, $C(R^{16})(R^{17})$ and NR^6;

E is selected from the group consisting of CH_2, O, S, and NR^7;

J is selected from the group consisting of O, S and NR^8;

T is selected from the group consisting of C(O) and $(CH_2)_b$ wherein b is an integer of from 0 to 3;

M is selected from the group consisting of $C(R^9)(R^{10})$ and $(CH_2)_u$, wherein u is an integer of from 0 to 3;

L is selected from the group consisting of O, NR^{11}, S, and $(CH_2)_n$ wherein n is an integer of 0 or 1;

X is selected from the group consisting of CO_2B, PO_3H_2, SO_3H, SO_2NH_2, SO_2NHCOR^{12}, OPO_3H_2, $C(O)NHC(O)R^{13}$, $C(O)NHSO_2R^{14}$, hydroxyl, tetrazolyl and hydrogen;

W is selected from the group consisting of C, CR^{15} and N;

B is selected from the group consisting of hydrogen, alkyl, alkenyl, alkynyl, hydroxyalkyl, haloalkyl, $-CF_3$, cycloalkyl, cycloalkenyl, cycloalkynyl, cycloalkylalkyl, aryl, heterocyclyl, alkylaryl, aralkenyl, aralkyl,

alkylheterocyclyl, heterocyclylalkyl and aryloxyalkyl; and

R^1, R^2, R^3, R^4, R^5, R^6, R^7, R^8, R^9, R^{10}, R^{11}, R^{12}, R^{13}, R^{14}, R^{15}, R^{16} and R^{17} at each occurrence are independently selected from the group consisting of hydrogen, halogen, alkyl, alkenyl, alkynyl, alkoxy, alkenoxy, alkynoxy, thioalkoxy, hydroxyalkyl, aliphatic acyl, $-CF_3$, $-CO_2H$, -SH, -CN, $-NO_2$, $-NH_2$, -OH, alkynylamino, alkoxycarbonyl, heterocycloyl, carboxy, $-N(C_1-C_3$ alkyl)$-C(O)(C_1-C_3$ alkyl), $-NHC(O)N(C_1-C_3$ alkyl)$C(O)NH(C_1-C_3$alkyl), $-NHC(O)NH(C_1-C_6$ alkyl), $-NHSO_2(C_1-C_3$ alkyl), $-NHSO_2$(aryl), alkoxyalkyl, alkylamino, alkenylamino, di(C_1-C_3)amino, $-C(O)O-(C_1-C_3)$alkyl, $-C(O)NH-(C_1-C_3)$alkyl, $-C(O)N(C_1-C_3$ alkyl)$_2$, -CH=NOH, $-PO_3H_2$, $-OPO_3H_2$, haloalkyl, alkoxyalkoxy, carboxaldehyde, carboxamide, cycloalkyl, cycloalkenyl, cycloalkynyl, cycloalkylalkyl, aryl, aroyl, aryloxy, arylamino, biaryl, thioaryl, diarylamino, heterocyclyl, alkylaryl, aralkenyl, aralkyl, alkylheterocyclyl, heterocyclylalkyl, sulfonyl, $-SO_2-(C_1-C_3$ alkyl), $-SO_3-(C_1-C_3$ alkyl), sulfonamido, carbamate, aryloxyalkyl and –C(O)NH(benzyl) groups;

wherein B, R^1, R^2, R^3, R^4, R^5, R^6, R^7, R^8, R^9, R^{10}, R^{11}, R^{12}, R^{13}, R^{14}, R^{15}, R^{16} and R^{17} are unsubstituted or substituted with at least one electron donating or electron withdrawing group;

wherein when L is NR^{11}, R^4 and R^{11} taken together may form a ring; and wherein when M is $C(R^9)(R^{10})$, R^9 and R^{10} taken together may form a ring;

and wherein when A is NR^6 and at least one Y is CR^1, R^1 and R^6 taken together may form a ring;

or a pharmaceutically acceptable salt thereof;

one or more other therapeutically active compounds and a pharmacologically acceptable diluent.

2. A composition of claim 1 wherein

A is NR^6;

E is NR^7;

J is O;

M is $C(R^9)(R^{10})$;

q is 4 or 5;

T is $(CH_2)_b$ wherein b is 0;

L is $(CH_2)_n$ wherein n is 0;

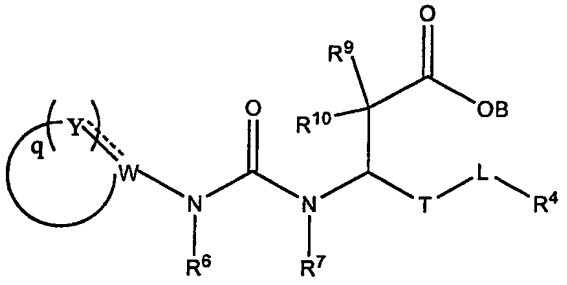
X is CO_2B;

W is C or CR^{15};

R^4 is selected from the group consisting of aryl, alkylaryl, aralkyl, heterocyclyl, alkylheterocyclyl and heterocyclylalkyl; and

R^6, R^7, R^9, R^{10} and R^{15} are independently selected from the group consisting of hydrogen and lower alkyl.

3. A pharmaceutical composition comprising a compound of the structure



wherein Y, at each occurrence, is independently selected from the group consisting of C(O), N, CR^1, $C(R^2)(R^3)$, NR^5, CH, O and S;

q is an integer of from 3 to 7;

T is selected from the group consisting of C(O) and $(CH_2)_b$ wherein b is an integer of 0 to 3;

L is selected from the group consisting of O, NR^{11}, S, and $(CH_2)_n$ wherein n is an integer of 0 or 1;

W is selected from the group consisting of C, CR^{15} and N;

B is selected from the group consisting of hydrogen, alkyl, alkenyl, alkynyl, hydroxyalkyl, haloalkyl, $-CF_3$, cycloalkyl, cycloalkenyl,

cycloalkynyl, cycloalkylalkyl, aryl, heterocyclyl, alkylaryl, aralkenyl, aralkyl, alkylheterocyclyl, heterocyclylalkyl and aryloxyalkyl; and

R^1, R^2, R^3, R^4, R^5, R^6, R^7, R^9, R^{10}, R^{11} and R^{15} are independently selected from the group consisting of hydrogen, halogen, alkyl, alkenyl, alkynyl, alkoxy, alkenoxy, alkynoxy, thioalkoxy, hydroxyalkyl, aliphatic acyl, $-CF_3$, $-CO_2H$, -SH, -CN, $-NO_2$, $-NH_2$, -OH, alkynylamino, alkoxycarbonyl, heterocycloyl, carboxy, $-N(C_1-C_3$ alkyl)$-C(O)(C_1-C_3$ alkyl), $-NHC(O)N(C_1-C_3$ alkyl)$C(O)NH(C_1-C_3$alkyl), $-NHC(O)NH(C_1-C_6$ alkyl), $-NHSO_2(C_1-C_3$ alkyl), $-NHSO_2$(aryl), alkoxyalkyl, alkylamino, alkenylamino, di(C_1-C_3)amino, $-C(O)O-(C_1-C_3)$alkyl, $-C(O)NH-(C_1-C_3)$alkyl, $-C(O)N(C_1-C_3$ alkyl$)_2$, -CH=NOH, $-PO_3H_2$, $-OPO_3H_2$, haloalkyl, alkoxyalkoxy, carboxaldehyde, carboxamide, cycloalkyl, cycloalkenyl, cycloalkynyl, cycloalkylalkyl, aryl, aroyl, aryloxy, arylamino, biaryl, thioaryl, diarylamino, heterocyclyl, alkylaryl, aralkenyl, aralkyl, alkylheterocyclyl, heterocyclylalkyl, sulfonyl, $-SO_2-(C_1-C_3$ alkyl), $-SO_3-(C_1-C_3$ alkyl), sulfonamido, carbamate, aryloxyalkyl and –C(O)NH(benzyl) groups;

wherein B, R^1, R^2, R^3, R^4, R^5, R^6, R^7, R^9, R^{10}, R^{11} and R^{15} are unsubstituted or substituted with at least one electron donating or electron withdrawing group;

wherein when L is NR^{11}, R^4 and R^{11} taken together may form a ring;

and wherein R^9 and R^{10} taken together may form a ring;

and wherein when at least one Y is CR^1, R^1 and R^6 taken together may form a ring;

or a pharmaceutically acceptable salt thereof;

one or more other therapeutically active compounds and a pharmacologically acceptable diluent.

4. A composition of claim 3 wherein

q is 4 or 5;

W is C or CR^{15};

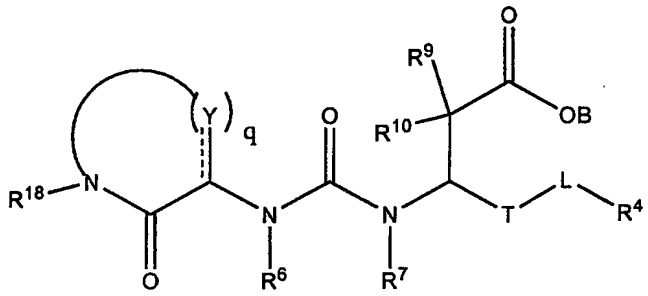
T is $(CH_2)_b$ wherein b is 0;

L is $(CH_2)_n$ wherein n is 0;

R^4 is selected from the group consisting of aryl, alkylaryl, aralkyl, heterocyclyl, alkylheterocyclyl and heterocyclylalkyl; and

R^6, R^7, R^9, R^{10} and R^{15} are independently selected from the group consisting of hydrogen and lower alkyl.

5. A pharmaceutical composition comprising a compound of the structure



wherein Y, at each occurrence, is independently selected from the group consisting of C(O), N, CR^1, $C(R^2)(R^3)$, NR^5, CH, O and S;

q is an integer of from 2 to 5;

T is selected from the group consisting of C(O) and $(CH_2)_b$ wherein b is an integer of 0 to 3;

L is selected from the group consisting of O, NR^{11}, S, and

$(CH_2)_n$ wherein n is an integer of 0 or 1;

R^5, R^6, R^7, R^{11} and R^{18} are each independently selected from the group consisting of alkyl, alkenyl, alkynyl, hydroxyalkyl, aliphatic acyl, alkynylamino, alkoxycarbonyl, heterocycloyl, -CH=NOH, haloalkyl, alkoxyalkoxy, carboxaldehyde, carboxamide, cycloalkyl, cycloalkenyl, cycloalkynyl, cycloalkylalkyl, aryl, aroyl, aryloxy, arylamino, biaryl, thioaryl, diarylamino, heterocyclyl, alkylaryl, aralkenyl, aralkyl, alkylheterocyclyl, heterocyclylalkyl, carbamate, aryloxyalkyl, hydrogen and –C(O)NH(benzyl) groups;

B is selected from the group consisting of hydrogen, alkyl, alkenyl, alkynyl, hydroxyalkyl, haloalkyl, $-CF_3$, cycloalkyl, cycloalkenyl, cycloalkynyl, cycloalkylalkyl, aryl, heterocyclyl, alkylaryl, aralkenyl, aralkyl, alkylheterocyclyl, heterocyclylalkyl and aryloxyalkyl; and

R^1, R^2, R^3, R^4, R^9 and R^{10} are independently selected from the group consisting of hydrogen, halogen, halkyl, alkenyl, alkynyl, alkoxy, alkenoxy, alkynoxy, thioalkoxy, hydroxyalkyl, aliphatic acyl, $-CF_3$, $-CO_2H$, -SH, -CN, $-NO_2$, $-NH_2$, -OH, alkynylamino, alkoxycarbonyl, heterocycloyl, carboxy, $-N(C_1$-C_3 alkyl)-$C(O)(C_1$-C_3 alkyl), $-NHC(O)N(C_1$-C_3 alkyl)$C(O)NH(C_1$-C_3alkyl), $-NHC(O)NH(C_1$-C_6 alkyl), $-NHSO_2(C_1$-C_3 alkyl), $-NHSO_2$(aryl), alkoxyalkyl,alkylamino, alkenylamino, di(C_1-C_3)amino, $-C(O)O-(C_1$-C_3)alkyl, $-C(O)NH-(C_1$-C_3)alkyl, $-C(O)N(C_1$-C_3 alkyl)$_2$, -CH=NOH, $-PO_3H_2$, $-OPO_3H_2$, haloalkyl, alkoxyalkoxy, carboxaldehyde, carboxamide, cycloalkyl, cycloalkenyl, cycloalkynyl, cycloalkylalkyl, aryl, aroyl, aryloxy, arylamino, biaryl, thioaryl, diarylamino, heterocyclyl, alkylaryl, aralkenyl, aralkyl, alkylheterocyclyl, heterocyclylalkyl, sulfonyl, $-SO_2-(C_1$-C_3 alkyl), $-SO_3-(C_1$-C_3 alkyl), sulfonamido, carbamate, aryloxyalkyl and –C(O)NH(benzyl) groups;

wherein B, R^1, R^2, R^3, R^4, R^5, R^6, R^7, R^9, R^{10}, R^{11} and R^{18} are unsubstituted or substituted with at least one electron donating or electron withdrawing group;

wherein when L is NR^{11}, R^4 and R^{11} taken together may form a ring;

and wherein R^9 and R^{10} taken together may form a ring;

and wherein when at least one Y is CR^1, R^1 and R^6 taken together may form a ring;

or a pharmaceutically acceptable salt thereof, one or more other therapeutically active compounds and a pharmacalogically acceptable diluent.

6. A composition of claim 5 wherein R^{18} is selected from the group consisting of hydrogen, alkyl, aryl, aralkyl, cycloalkyl, alkylheterocyclyl, heterocyclylalkyl and heterocyclyl;

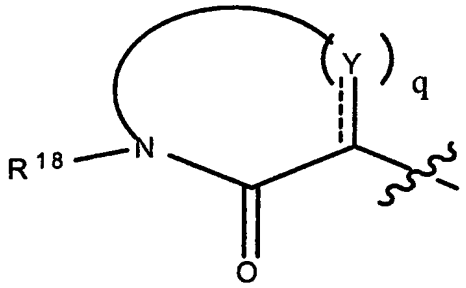
T is $(CH_2)_b$ wherein b is 0;

L is $(CH_2)_n$ wherein n is 0;

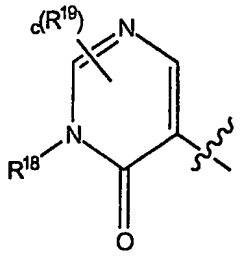
Y is selected from the group consisting of CR^1 and $C(R^2)(R^3)$ and

q is 2 or 3.

7. A composition of claim 5 wherein



is selected from the group consisting of



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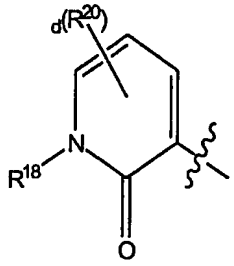
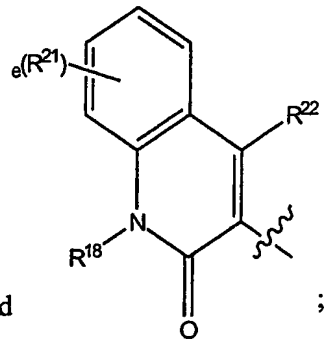


and



;

wherein R^{19}, R^{20}, R^{21} and R^{28} at each occurrence are independently selected from the group consisting of halogen, alkyl, alkenyl, alkynyl, alkoxy, alkenoxy, alkynoxy, thioalkoxy, hydroxyalkyl, aliphatic acyl, $-CF_3$, -OH, $-CO_2H$, -SH, -CN, $-NO_2$, $-NH_2$, alkynylamino, alkoxycarbonyl, heterocycloyl, carboxy, $-N(C_1-C_3$ alkyl)-$C(O)(C_1-C_3$ alkyl), $-NHC(O)N(C_1-C_3$ alkyl)$C(O)NH(C_1-C_3$alkyl), $-NHC(O)NH(C_1-C_6$ alkyl), $-NHSO_2(C_1-C_3$ alkyl), $-NHSO_2$(aryl), alkoxyalkyl, alkylamino, alkenylamino, di(C_1-C_3)amino, $-C(O)O-(C_1-C_3)$alkyl, $-C(O)NH-(C_1-C_3)$alkyl, $-C(O)N(C_1-C_3$ alkyl)$_2$, -CH=NOH, $-PO_3H_2$, $-OPO_3H_2$, haloalkyl, alkoxyalkoxy, carboxaldehyde, carboxamide, cycloalkyl, cycloalkenyl, cycloalkynyl, cycloalkylalkyl, aryl, aroyl, aryloxy, arylamino, biaryl, thioaryl, diarylamino, heterocyclyl, alkylaryl, aralkenyl, aralkyl, alkylheterocyclyl, heterocyclylalkyl, sulfonyl, $-SO_2-(C_1-C_3$ alkyl), $-SO_3-(C_1-C_3$ alkyl), sulfonamido, carbamate, aryloxyalkyl and –C(O)NH(benzyl) groups;

R^{18} is selected from the group consisting of alkyl, alkenyl, alkynyl, hydroxyalkyl, aliphatic acyl, alkynylamino, alkoxycarbonyl, heterocycloyl, -CH=NOH, haloalkyl, alkoxyalkoxy, carboxaldehyde, carboxamide, cycloalkyl, cycloalkenyl, cycloalkynyl, cycloalkylalkyl, aryl, aroyl, aryloxy, arylamino, biaryl, thioaryl, diarylamino, heterocyclyl, alkylaryl, aralkenyl, aralkyl, alkylheterocyclyl, heterocyclylalkyl, carbamate, aryloxyalkyl, hydrogen and –C(O)NH(benzyl) groups;

R^{22} is selected from the group consisting of hydrogen, halogen, alkyl, alkenyl, alkynyl, alkoxy, alkenoxy, alkynoxy, thioalkoxy, hydroxyalkyl, aliphatic acyl, $-CF_3$, $-CO_2H$, -SH, -CN, $-NO_2$, $-NH_2$, -OH, alkynylamino, alkoxycarbonyl, heterocycloyl, carboxy, $-N(C_1-C_3$ alkyl)-$C(O)(C_1-C_3$ alkyl), $-NHC(O)N(C_1-C_3$ alkyl)$C(O)NH(C_1-C_3$alkyl), $-NHC(O)NH(C_1-C_6$ alkyl), $-NHSO_2(C_1-C_3$ alkyl), $-NHSO_2$(aryl), alkoxyalkyl, alkylamino, alkenylamino, di(C_1-C_3)amino, $-C(O)O-(C_1-C_3)$alkyl,

-C(O)NH-(C_1-C_3)alkyl, -C(O)N(C_1-C_3 alkyl)$_2$, -CH=NOH, -PO_3H_2, -OPO_3H_2, haloalkyl, alkoxyalkoxy, carboxaldehyde, carboxamide, cycloalkyl, cycloalkenyl, cycloalkynyl, cycloalkylalkyl, aryl, aroyl, aryloxy, arylamino, biaryl, thioaryl, diarylamino, heterocyclyl, alkylaryl, aralkenyl, aralkyl, alkylheterocyclyl, heterocyclylalkyl, sulfonyl, -SO_2-(C_1-C_3 alkyl), -SO_3-(C_1-C_3 alkyl), sulfonamido, carbamate, aryloxyalkyl and –C(O)NH(benzyl) groups;

c is an integer of zero to two;

d is an integer of zero to three;

e is an integer of zero to four; and

i is an integer of zero to two.

8. The composition of claim 5 wherein R^{18} is aralkyl;

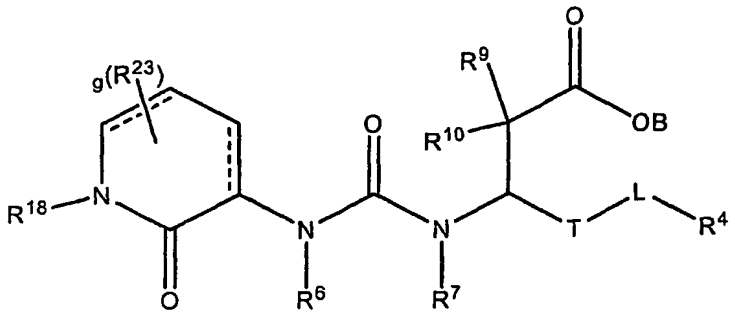
R^4 is aryl;

T is $(CH_2)_b$ where b is zero;

L is $(CH_2)_n$ where n is zero; and,

B, R^6, R^7, R^9 and R^{10} are each independently hydrogen.

9. A pharmaceutical composition comprising a compound of the structure



wherein T is selected from the group consisting of C(O) and $(CH_2)_b$ wherein b is an integer of from 0 to 3;

L is selected from the group consisting of O, NR^{11}, S, and

$(CH_2)_n$ wherein n is an integer of 0 or 1;

g is an integer of from 0 to 7;

B is selected from the group consisting of hydrogen, alkyl, alkenyl, alkynyl, hydroxyalkyl, haloalkyl, $-CF_3$, cycloalkyl, cycloalkenyl, cycloalkynyl, cycloalkylalkyl, aryl, heterocyclyl, alkylaryl, aralkenyl, aralkyl, alkylheterocyclyl, heterocyclylalkyl and aryloxyalkyl;

R^4, R^9, R^{10} and R^{23} at each occurrence are independently selected from the group consisting of hydrogen, halogen, alkyl, alkenyl, alkynyl, alkoxy, alkenoxy, alkynoxy, thioalkoxy, hydroxyalkyl, aliphatic acyl, $-CF_3$, $-CO_2H$, -SH, -CN, $-NO_2$, $-NH_2$, -OH, alkynylamino, alkoxycarbonyl, heterocycloyl, carboxy, $-N(C_1-C_3$ alkyl)$-C(O)(C_1-C_3$ alkyl), $-NHC(O)N(C_1-C_3$ alkyl)$C(O)NH(C_1-C_3$alkyl), $-NHC(O)NH(C_1-C_6$ alkyl), $-NHSO_2(C_1-C_3$ alkyl), $-NHSO_2$(aryl), alkoxyalkyl,alkylamino, alkenylamino, di(C_1-C_3)amino, $-C(O)O-(C_1-C_3)$alkyl, $-C(O)NH-(C_1-C_3)$alkyl, $-C(O)N(C_1-C_3$ alkyl$)_2$, -CH=NOH, $-PO_3H_2$, $-OPO_3H_2$, haloalkyl, alkoxyalkoxy, carboxaldehyde, carboxamide, cycloalkyl, cycloalkenyl, cycloalkynyl, cycloalkylalkyl, aryl, aroyl, aryloxy, arylamino, biaryl, thioaryl, diarylamino, heterocyclyl, alkylaryl, aralkenyl, aralkyl, alkylheterocyclyl, heterocyclylalkyl, sulfonyl, $-SO_2-(C_1-C_3$ alkyl), $-SO_3-(C_1-C_3$ alkyl), sulfonamido, carbamate, aryloxyalkyl and –C(O)NH(benzyl) groups;

R^6, R^7, R^{11} and R^{18} are each independently selected from the group consisting of alkyl, alkenyl, alkynyl, hydroxyalkyl, aliphatic acyl, alkynylamino, alkoxycarbonyl, heterocycloyl, -CH=NOH, haloalkyl, alkoxyalkoxy, carboxaldehyde, carboxamide, cycloalkyl, cycloalkenyl, cycloalkynyl, cycloalkylalkyl, aryl, aroyl, aryloxy, arylamino, biaryl, thioaryl, diarylamino, heterocyclyl, alkylaryl, aralkenyl, aralkyl, alkylheterocyclyl, heterocyclylalkyl, carbamate, aryloxyalkyl, hydrogen and –C(O)NH(benzyl) groups;

wherein B, R^4, R^6, R^7, R^9, R^{10}, R^{11}, R^{18} and R^{23} are unsubstituted or substituted with at least one electron donating or electron withdrawing group;

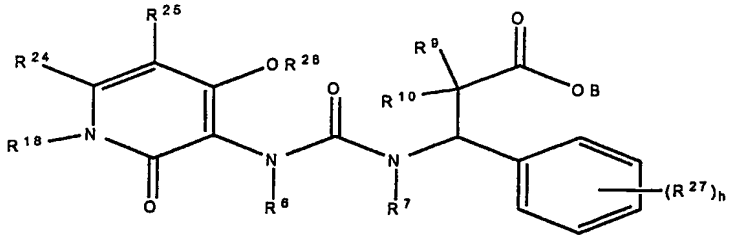
wherein when L is NR^{11}, R^4 and R^{11} taken together may form a ring;

and wherein R^9 and R^{10} taken together may form a ring;

or a pharmaceutically acceptable salt thereof;

one or more other therapeutically active compounds and a pharmacologically acceptable diluent.

10. A pharmaceutical composition comprising a compound of the structure



wherein h is an integer of zero to five;

B is selected from the group consisting of hydrogen, alkyl, alkenyl, alkynyl, hydroxyalkyl, haloalkyl, $-CF_3$, cycloalkyl, cycloalkenyl, cycloalkynyl, cycloalkylalkyl, aryl, heterocyclyl, alkylaryl, aralkenyl, aralkyl, alkylheterocyclyl, heterocyclylalkyl and aryloxyalkyl;

R^9, R^{10}, R^{24} and R^{25} are each independently selected from the group consisting of hydrogen, halogen, alkyl, alkenyl, alkynyl, alkoxy, alkenoxy, alkynoxy, thioalkoxy, hydroxyalkyl, aliphatic acyl, $-CF_3$, $-CO_2H$, -SH, -CN, $-NO_2$, $-NH_2$, -OH, alkynylamino, alkoxycarbonyl, heterocycloyl, carboxy, $-N(C_1\text{-}C_3$ alkyl)$-C(O)(C_1\text{-}C_3$ alkyl), $-NHC(O)N(C_1\text{-}C_3$ alkyl)$C(O)NH(C_1\text{-}C_3$alkyl), $-NHC(O)NH(C_1\text{-}C_6$ alkyl), $-NHSO_2(C_1\text{-}C_3$ alkyl), $-NHSO_2$(aryl), alkoxyalkyl, alkylamino, alkenylamino, di$(C_1\text{-}C_3)$amino, $-C(O)O\text{-}(C_1\text{-}C_3)$alkyl, $-C(O)NH\text{-}(C_1\text{-}C_3)$alkyl, $-C(O)N(C_1\text{-}C_3$ alkyl$)_2$, -CH=NOH, $-PO_3H_2$, $-OPO_3H_2$, haloalkyl, alkoxyalkoxy, carboxaldehyde, carboxamide, cycloalkyl, cycloalkenyl, cycloalkynyl, cycloalkylalkyl, aryl, aroyl, aryloxy, arylamino, biaryl, thioaryl, diarylamino, heterocyclyl, alkylaryl,

aralkenyl, aralkyl, alkylheterocyclyl, heterocyclylalkyl, sulfonyl, -SO_2-(C_1-C_3 alkyl), -SO_3-(C_1-C_3 alkyl), sulfonamido, carbamate, aryloxyalkyl and –C(O)NH(benzyl) groups;

R^{27}, at each occurrence, is independently selected from the group consisting of halogen, hydroxyl, alkyl, alkenyl, alkynyl, alkoxy, alkenoxy, alkynoxy, thioalkoxy, hydroxyalkyl, aliphatic acyl, -CF_3, -CO_2H, -SH, -CN, -NO_2, -NH_2, alkynylamino, alkoxycarbonyl, heterocycloyl, carboxy, -N(C_1-C_3 alkyl)-C(O)(C_1-C_3 alkyl), -NHC(O)N(C_1-C_3 alkyl)C(O)NH(C_1-C_3alkyl), -NHC(O)NH(C_1-C_6 alkyl), -$NHSO_2$(C_1-C_3 alkyl), -$NHSO_2$(aryl), -N(C_1-C_3alkyl)SO_2(C_1-C_3alkyl), -N(C_1-C_3alkyl)SO_2(aryl), alkoxyalkyl, alkylamino, alkenylamino, di(C_1-C_3)amino, -C(O)O-(C_1-C_3)alkyl, -C(O)NH-(C_1-C_3)alkyl, -C(O)N(C_1-C_3 alkyl)$_2$, -CH=NOH, -PO_3H_2, -OPO_3H_2, haloalkyl, alkoxyalkoxy, carboxaldehyde, carboxamide, cycloalkyl, cycloalkenyl, cycloalkynyl, cycloalkylalkyl, aryl, aroyl, aryloxy, arylamino, biaryl, thioaryl, diarylamino, heterocyclyl, alkylaryl, aralkenyl, aralkyl, alkylheterocyclyl, heterocyclylalkyl, sulfonyl, -SO_2-(C_1-C_3 alkyl), -SO_3-(C_1-C_3 alkyl), sulfonamido, carbamate, aryloxyalkyl and –C(O)NH(benzyl) groups;

R^6, R^7 and R^{18} are each independently selected from the group consisting of alkyl, alkenyl, alkynyl, hydroxyalkyl, aliphatic acyl, alkynylamino, alkoxycarbonyl, heterocycloyl, -CH=NOH, haloalkyl, alkoxyalkoxy, carboxaldehyde, carboxamide, cycloalkyl, cycloalkenyl, cycloalkynyl, cycloalkylalkyl, aryl, aroyl, aryloxy, arylamino, biaryl, thioaryl, diarylamino, heterocyclyl, alkylaryl, aralkenyl, aralkyl, alkylheterocyclyl, heterocyclylalkyl, carbamate, aryloxyalkyl, hydrogen and –C(O)NH(benzyl) groups; and,

R^{26} is selected from the group consisting of hydrogen, alkyl, alkenyl, alkynyl, hydroxyalkyl, aliphatic acyl, -CF_3, alkoxycarbonyl, heterocycloyl, carboxy, -C(O)O-(C_1-C_3)alkyl, -C(O)NH-(C_1-C_3)alkyl, -C(O)N(C_1-C_3 alkyl)$_2$, -PO_3H_2, haloalkyl, carboxamide, cycloalkyl, cycloalkenyl, cycloalkynyl, cycloalkylalkyl, aryl, aroyl, biaryl, heterocyclyl, alkylaryl, aralkenyl,

aralkyl, alkylheterocyclyl, heterocyclylalkyl, sulfonyl, -SO_2-(C_1-C_3 alkyl), sulfonamido, aryloxyalkyl and –C(O)NH(benzyl) groups;

wherein B, R^6, R^7, R^9, R^{10}, R^{18}, R^{24}, R^{25}, R^{26} and R^{27} are unsubstituted or substituted with at least one electron donating or electron withdrawing group;

wherein R^{18} and R^{24} taken together may form a ring;

R^{24} and R^{25} taken together may form a ring;

R^{25} and R^{26} taken together may form a ring;

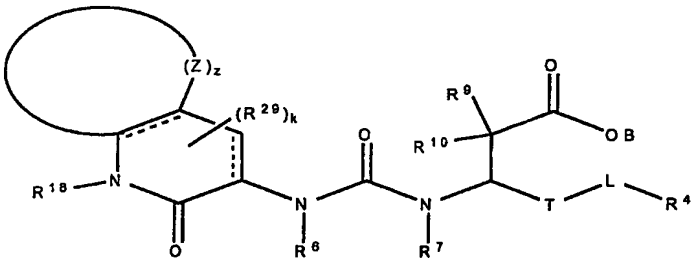
and wherein R^9 and R^{10} taken together may form a ring;

or a pharmaceutically acceptable salt thereof;

one or more other therapeutically active compounds and a pharmacologically acceptable diluent.

11. The composition of claim 10 wherein B, R^6, R^7, R^9, R^{10}, R^{24}, R^{25} and R^{26} are each independently hydrogen and R^{18} is substituted or unsubstituted aralkyl.

12. A pharmaceutical composition comprising a compound of the structure



wherein Z, at each occurrence, is independently selected from the group consisting of C(O), N, CR^{30}, $C(R^{31})(R^{32})$, NR^{33}, CH, O and S;

z is an integer of from 3 to 6;

k is an integer of from 0 to 5;

T is selected from the group consisting of C(O) and $(CH_2)_b$ wherein b is an integer of from 0 to 3;

L is selected from the group consisting of O, NR^{11}, S, and

$(CH_2)_n$ wherein n is an integer of 0 or 1;

R^6, R^7, R^{11}, R^{18} and R^{33} are each independently selected from the group consisting of alkyl, alkenyl, alkynyl, hydroxyalkyl, aliphatic acyl, alkynylamino, alkoxycarbonyl, heterocycloyl, -CH=NOH, haloalkyl, alkoxyalkoxy, carboxaldehyde, carboxamide, cycloalkyl, cycloalkenyl, cycloalkynyl, cycloalkylalkyl, aryl, aroyl, aryloxy, arylamino, biaryl, thioaryl, diarylamino, heterocyclyl, alkylaryl, aralkenyl, aralkyl, alkylheterocyclyl, heterocyclylalkyl, carbamate, aryloxyalkyl, hydrogen and –C(O)NH(benzyl) groups;

B is selected from the group consisting of hydrogen, alkyl, alkenyl, alkynyl, hydroxyalkyl, haloalkyl, $-CF_3$, cycloalkyl, cycloalkenyl, cycloalkynyl, cycloalkylalkyl, aryl, heterocyclyl, alkylaryl, aralkenyl, aralkyl, alkylheterocyclyl, heterocyclylalkyl and aryloxyalkyl;

R^4, R^9, R^{10}, R^{30}, R^{31} and R^{32} at each occurrence are independently selected from the group consisting of hydrogen, halogen, alkyl, alkenyl, alkynyl, alkoxy, alkenoxy, alkynoxy, thioalkoxy, hydroxyalkyl, aliphatic acyl, $-CF_3$, $-CO_2H$, -SH, -OH, -CN, $-NO_2$, $-NH_2$, alkynylamino, alkoxycarbonyl, heterocycloyl, carboxy, $-N(C_1-C_3$ alkyl)$-C(O)(C_1-C_3$ alkyl), $-NHC(O)N(C_1-C_3$ alkyl)$C(O)NH(C_1-C_3$alkyl), $-NHC(O)NH(C_1-C_6$ alkyl), $-NHSO_2(C_1-C_3$ alkyl), $-NHSO_2$(aryl), alkoxyalkyl, alkylamino, alkenylamino, di(C_1-C_3)amino, $-C(O)O-(C_1-C_3)$alkyl, $-C(O)NH-(C_1-C_3)$alkyl, $-C(O)N(C_1-C_3$ alkyl$)_2$, -CH=NOH, $-PO_3H_2$, $-OPO_3H_2$, haloalkyl, alkoxyalkoxy, carboxaldehyde, carboxamide, cycloalkyl, cycloalkenyl, cycloalkynyl, cycloalkylalkyl, aryl, aroyl, aryloxy, arylamino, biaryl, thioaryl, diarylamino, heterocyclyl, alkylaryl, aralkenyl, aralkyl, alkylheterocyclyl, heterocyclylalkyl, sulfonyl, $-SO_2-(C_1-C_3$ alkyl), $-SO_3-(C_1-C_3$ alkyl), sulfonamido, carbamate, aryloxyalkyl and –C(O)NH(benzyl) groups; and

R^{29}, at each occurrence, is independently selected from the group consisting of halogen, alkyl, alkenyl, alkynyl, alkoxy, alkenoxy, alkynoxy, thioalkoxy, hydroxyalkyl, aliphatic acyl, $-CF_3$, $-CO_2H$, -SH, -CN, $-NO_2$, $-NH_2$, -OH, alkynylamino, alkoxycarbonyl, heterocycloyl, carboxy, $-N(C_1-C_3$ alkyl)-

$C(O)(C_1\text{-}C_3$ alkyl), $-NHC(O)N(C_1\text{-}C_3$ alkyl)$C(O)NH(C_1\text{-}C_3$alkyl), $-NHC(O)NH(C_1\text{-}C_6$ alkyl), $-NHSO_2(C_1\text{-}C_3$ alkyl), $-NHSO_2$(aryl), alkoxyalkyl, alkylamino, alkenylamino, di$(C_1\text{-}C_3)$amino, $-C(O)O\text{-}(C_1\text{-}C_3)$alkyl, $-C(O)NH\text{-}(C_1\text{-}C_3)$alkyl, $-C(O)N(C_1\text{-}C_3$ alkyl$)_2$, -CH=NOH, $-PO_3H_2$, $-OPO_3H_2$, haloalkyl, alkoxyalkoxy, carboxaldehyde, carboxamide, cycloalkyl, cycloalkenyl, cycloalkynyl, cycloalkylalkyl, aryl, aroyl, aryloxy, arylamino, biaryl, thioaryl, diarylamino, heterocyclyl, alkylaryl, aralkenyl, aralkyl, alkylheterocyclyl, heterocyclylalkyl, sulfonyl, $-SO_2\text{-}(C_1\text{-}C_3$ alkyl), $-SO_3\text{-}(C_1\text{-}C_3$ alkyl), sulfonamido, carbamate, aryloxyalkyl and –C(O)NH(benzyl) groups;

wherein B, R^4, R^5, R^6, R^7, R^9, R^{10}, R^{11}, R^{18}, R^{29}, R^{30}, R^{31}, R^{32} and R^{33} are unsubstituted or substituted with at least one electron donating or electron withdrawing group;

wherein when L is NR^{11}, R^4 and R^{11} taken together may form a ring;

and wherein R^9 and R^{10} taken together may form a ring;

or a pharmaceutically acceptable salt thereof;

one or more other therapeutically active compounds and a pharmacologically acceptable diluent.

13. The composition of claim 12 wherein z is three or four.

14. The composition of claim 1 where the compound of structure (I) is (3S)-3-[({[1-(2-chlorobenzyl)-4-hydroxy-5-methyl-2-oxo-1,2-dihydropyridin-3-yl]amino}carbonyl)amino]-3-(4-methylphenyl)propanoic acid and pharmaceutically acceptable salts thereof.

15. The composition of claim 1 where the compound of structure (I) is (3S)-3-[({[1-(2-chlorobenzyl)-4-hydroxy-2-oxo-2,5,6,7-tetrahydro-1H-cyclopenta[b]pyridin-3-l]amino}carbonyl)amino]-3-(4-methylphenyl)propanoic acid and pharmaceutically acceptable salts thereof.

16. The composition of claim 1 where the compound of structure (I) is (3S)-3-[({[1-(2-

chlorobenzyl)-4-hydroxy-5-methyl-2-oxo-1,2-dihydropyridin-3-yl]amino}carbonyl)amino]-3-[3-(diethylamino)phenyl]propanoic acid and pharmaceutically acceptable salts thereof.

17. The composition of claim 1 where the compound of structure (I) is (3S)-3-[({[1-(2-chlorobenzyl)-4-hydroxy-2-oxo-2,5,6,7-tetrahydro-1H-cyclopenta[b]pyridin-3-yl] amino}carbonyl)amino]-3-(3-isopropylphenyl)propanoic acid and pharmaceutically acceptable salts thereof.

18. The composition of claim 1 where the compound of structure (I) is (3S)-3-[({[1-(2-chlorobenzyl)-4-hydroxy-2-oxo-1,2-dihydro-1,8-napthyridin-3-yl]amino} carbonyl)amino-3-(4-methylphenyl)propanoic acid and pharmaceutically acceptable salts thereof.

19. The composition of claim 1 where the other therapeutically active compounds are selected from the group consisting of IL-5 antagonists, CCR-3 antagonists, corticosteroids, antihistamines, Leukotrine antagonists, COX-I and COX-II inhibitors, mast cell stabilizers, anti IL-5 and anti IgE antibodies, IL-5 synthesis and release inhibitors, TNF-α inhibitors, p38 MAP kinase inhibitors, tryptase inhibitors, anticytokine/antichemokine agents, vaccines, cromolyn, selectin antagonists, PDE 4 inhibitors, β-agonists, muscarininc antagonists and immunosuppressives, CD20 antagonists and syk tyrosine kinase inhibitors.

20. A method for treating an inflammatory disease in a mammal comprising administering to said mammal a therapeutically effective amount of a composition of claim 1.

21. The method of claim 20 wherein the inflammatory disease is selected from psoriasis, asthma, atherosclerosis, multiple sclerosis, Guillan-Barr Syndrome, rheumatoid arthritis, inflammatory bowel disease and reperfusion injury.

22. A method for treating an inflammatory disease in a mammal comprising administering to said mammal a therapeutically effective amount of a combination of a compound of structure (I) in claim 1 and an effective amount of one or more other therapeutic compounds.

23. The method of claim 22 wherein the inflammatory disease is selected from psoriasis, asthma, atherosclerosis, multiple sclerosis, Guillan-Barr Syndrome, rheumatoid arthritis, inflammatory bowel disease and reperfusion injury.

24. The composition of claim 19 wherein the compound of structure (I) is selected from the group consisting of (3S)-3-[({[1-(2-chlorobenzyl)-4-hydroxy-5-methyl-2-oxo-1,2-dihydropyridin-3-yl]amino}carbonyl)amino]-3-(4-methylphenyl)propanoic acid; (3S)-3-[({[1-(2-chlorobenzyl)-4-hydroxy-2-oxo-2,5,6,7-tetrahydro-1H-cyclopenta [b]pyridin-3-l]amino}carbonyl)amino]-3-(4-methylphenyl)propanoic acid; (3S)-3-[({[1-(2-chlorobenzyl)-4-hydroxy-5-methyl-2-oxo-1,2-dihydropyridin-3-yl]amino}carbonyl)amino]-3-[3-(diethylamino)phenyl]propanoic acid; (3S)-3-[({[1-(2-chlorobenzyl)-4-hydroxy-2-oxo-2,5,6,7-tetrahydro-1H-cyclopenta[b]pyridin-3-yl] amino}carbonyl)amino]-3-(3-isopropylphenyl)propanoic acid; and (3S)-3-[({[1-(2-chlorobenzyl)-4-hydroxy-2-oxo-1,2-dihydro-1,8-napthyridin-3-yl]amino} carbonyl) amino-3-(4-methylphenyl)propanoic acid and pharmaceutically acceptable salts thereof.

25. The method of claim 20 wherein the compound of structure (I) is selected from the group consisting of (3S)-3-[({[1-(2-chlorobenzyl)-4-hydroxy-5-methyl-2-oxo-1,2-dihydropyridin-3-yl]amino}carbonyl)amino]-3-(4-methylphenyl)propanoic acid; (3S)-3-[({[1-(2-chlorobenzyl)-4-hydroxy-2-oxo-2,5,6,7-tetrahydro-1H-cyclopenta [b]pyridin-3-l]amino}carbonyl)amino]-3-(4-methylphenyl)propanoic acid; (3S)-3-[({[1-(2-chlorobenzyl)-4-hydroxy-5-methyl-2-oxo-1,2-dihydropyridin-3-yl]amino}carbonyl)amino]-3-[3-(diethylamino)phenyl]propanoic acid; (3S)-3-[({[1-(2-chlorobenzyl)-4-hydroxy-2-oxo-2,5,6,7-tetrahydro-1H-cyclopenta[b]pyridin-3-yl] amino}carbonyl)amino]-3-(3-isopropylphenyl)propanoic acid; and (3S)-3-[({[1-(2-chlorobenzyl)-4-hydroxy-2-oxo-1,2-dihydro-1,8-napthyridin-3-yl]amino} carbonyl) amino-3-(4-methylphenyl)propanoic acid and pharmaceutically acceptable salts thereof.

26. A kit comprising in a single package, one container comprising a compound that inhibits binding of an $\alpha_4\beta_1$ integrin to its receptors as set forth in structure (I) in claim 1 in a

pharmaceutically acceptable carrier and one or more separate containers comprising other therapeutic compounds in pharmaceutically acceptable carriers, with the compound that inhibits binding of $\alpha_4\beta_1$ integrin to its receptors and the other therapeutic compounds being present in amounts such that the combination is effective to treat disease states mediated by $\alpha_4\beta_1$ integrin binding.